| AD | ) |  |  |  |
|----|---|--|--|--|
|    |   |  |  |  |

Award Number: W81XWH-11-2-0040

TITLE: AltitudeOmics: The Basic Biology of Human Acclimatization to High Altitude

PRINCIPAL INVESTIGATOR: Dr. Robert Roach

CONTRACTING ORGANIZATION: The University of Colorado Aurora, CO 80045

REPORT DATE: January 2012

TYPE OF REPORT: Annual

PREPARED FOR: U.S. Army Medical Research and Materiel Command Fort Detrick, Maryland 21702-5012

DISTRIBUTION STATEMENT: Approved for public release; distribution unlimited

The views, opinions and/or findings contained in this report are those of the author(s) and should not be construed as an official Department of the Army position, policy or decision unless so designated by other documentation.

| R                                     | EPORT DO  |   | Form Approved<br>OMB No. 0704-0188                                      |                         |   |
|---------------------------------------|---|---|---|-------------------------|---|
| Public reporting burden for this      | collection of information is est                            | timated to average 1 hour per re  | esponse, including the time for revi                                    |                         | ching existing data sources, gathering and maintaining the  |
| this burden to Department of De       | efense, Washington Headqua<br>aware that notwithstanding an | rters Services, Directorate for Ir<br>ny other provision of law, no per | formation Operations and Reports<br>son shall be subject to any penalty | (0704-0188), 1215 Jeffe | ollection of information, including suggestions for reducing<br>erson Davis Highway, Suite 1204, Arlington, VA 22202-<br>n a collection of information if it does not display a currently |
| 1. REPORT DATE (DD                    |   | 2. REPORT TYPE  |   |                         | DATES COVERED (From - To)   |
| 01-01-2012<br>4. TITLE AND SUBTITE    | <u> </u><br>L <b>F</b>                                      | Annual  |   |                         | IAN 2011 - 31 DEC 2011<br>CONTRACT NUMBER   |
|                                       | <del></del>   | Juman Acclimatiza   | tion to High Altitude   | Jul.                    | CONTRACT NOMBER   |
| AilitudeOffics. The                   | Dasic biology of i  | iuman Acciimatiza   | tion to riigh Attitude  | 5b.                     | GRANT NUMBER  |
|                                       |   |   |   |                         | V81XWH-11-2-0040  |
|                                       |   |   |   | 5c.                     | PROGRAM ELEMENT NUMBER  |
| 6. AUTHOR(S)                          |   |   |   | 5d.                     | PROJECT NUMBER  |
| Dr. Robert Roach                      |   |   |   | - Fa                    | TACK NUMBER   |
|                                       |   |   |   | 50.                     | TASK NUMBER   |
| E-Mail: robert.road                   | ch@ucdenver.edu   |   |   | 5f. \                   | WORK UNIT NUMBER  |
| 7. PERFORMING ORG                     | ANIZATION NAME(S)   | ) AND ADDRESS(ES)   |   |                         | PERFORMING ORGANIZATION REPORT  |
| The University of C                   | olorado   |   |   | "                       | IUMBER  |
| Aurora, CO 80045                      |   |   |   |                         |   |
|                                       |   |   |   |                         |   |
|                                       |   |   |   |                         |   |
| 9. SPONSORING / MO                    | NITORING AGENCY   | NAME(S) AND ADDRE   | SS(ES)  | 10.                     | SPONSOR/MONITOR'S ACRONYM(S)  |
| U.S. Army Medical                     |   |   |   |                         |   |
| Fort Detrick, Maryla                  | and 21702-5012  |   |   |                         |   |
|                                       |   |   |   |                         | SPONSOR/MONITOR'S REPORT<br>NUMBER(S)   |
| 12. DISTRIBUTION / A                  | _   |   |   | l                       |   |
| Approved for Public 13. SUPPLEMENTARY |   | ution Unlimited   |   |                         |   |
| 14. ABSTRACT                          |   |   |   |                         |   |
| 14. ADSTRACT                          |   |   |   |                         |   |
|                                       |   |   |   |                         |   |
| Progress is being m                   | nade on all aspect  | s of this projects Th   | ne 65 nage protocol a   | and 25 nage in          | formed consent documents have   |
|                                       |   |   |   |                         | and University of Oregon. Those   |
|                                       | •   |   |   |                         | ned and are currently being tested.   |
|                                       | •   |   | -   | _                       | atory first to Oregon for pre-testing   |
|                                       |   |   |   |                         | expect to meet all future milestones  |
| and to successfully                   | complete the field  | I portion of this stud  | dy in September 2012  | 2.                      |   |
|                                       |   |   |   |                         |   |
|                                       |   |   |   |                         |   |
|                                       |   |   |   |                         |   |
|                                       |   |   |   |                         |   |
| 15. SUBJECT TERMS                     |   |   |   |                         |   |
| Hypoxia, High Altitu                  | de Acclimatization  | n, systems biology,   | epigenetics, gene ex  | pression, prote         | eomics, metabolomics  |
| 16. SECURITY CLASS                    | IFICATION OF:   |   | 17. LIMITATION  | 18. NUMBER              | 19a. NAME OF RESPONSIBLE PERSON   |
| OLOUNIT OLAGO                         |   |   | OF ABSTRACT   | OF PAGES                | USAMRMC   |
| a. REPORT                             | b. ABSTRACT   | c. THIS PAGE  |   | 4                       | 19b. TELEPHONE NUMBER (include area code)   |

# **Table of Contents**

|                              | Page |
|------------------------------|------|
|                              |      |
|                              |      |
|                              |      |
|                              |      |
| Introduction                 | 4    |
| Body                         | 4    |
| Key Research Accomplishments | 4    |
| Reportable Outcomes          | 4    |
| Conclusion                   | 4    |
| References                   | 4    |
| Appendices                   | 4    |

## INTRODUCTION:

The goal of this project is to advance high-altitude medical research by discovering the basic molecular mechanisms of acclimatization and de-acclimatization that protect soldiers from high-altitude illness.

## **BODY:**

In this first year we have accomplished most of the tasks outlined in the statement of work for months 1-12.

- We have obtained IRB approvals from University of Colorado Denver and the University of Oregon where our subjects will be recruited. Our submission is currently being reviewed by DOD-HRPO.
- We have also started the logistical preparations for this complex field study. We have made a first preparatory
  visit to Bolivia to inspect the status of the laboratory and housing we will use, and have started the process for
  necessary approvals and improvements.
- We have also begun to practice the data analysis pipeline. We have developed an optimized plan for all AltitudeOmics data analysis. Given the fact that we will generate more than 3.5 million data points for gene expression only, this has been a critical accomplishment.
- We have drafted the first two papers from this study.

According to our milestones we should have started screening and recruiting subjects in Q4. However, since we have not received HRPO approval, we have not been able to start this process. We are prepared to start the day we receive approval. Like the AMS Prediction study, we really are able to move quickly once approvals are in place. If the HRPO process runs smoothly, we should catch up by the end of the 2<sup>nd</sup> quarter 2012.

# **KEY RESEARCH ACCOMPLISHMENTS:**

The goal of this stage of the planning is put all procedures into place that will allow us to complete this multi-faceted, multi-investigator, multi-national project in between Phase I and Phase II of our other funded TATRC project: AMS Prediction. I've led or been part of dozens of research expeditions in my career. AltitudeOmics is more organized, with a better chance of scientific success than any of our previous expeditions. Our emphasis on pre-planning analysis and interpretation leaves room for innovation and new discoveries, but also will speed up the production of results from the study.

The key papers that will come out of the physiology portion of the study will be published first, and then in a few months after returning form Bolivia the first round of OMICS papers will come out. Toward the end of the project performance period the papers integrating the results form the standalone physiology and OMICS papers will be integrated into several new reports that should surpass anything previously done in this filed in terms of integrating physiology and the cellular and molecular biology of acclimatization.

# **REPORTABLE OUTCOMES:**

Having just laid the groundwork for this study, which we hope to conduct during the summer of 2012, we do not have any reportable outcomes yet.

#### **CONCLUSION:**

There are no conclusions to be drawn yet from the work performed to date.